

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application.

Listing of Claims:

1. (Currently Amended) ~~A microfluidic device~~ An analytical system comprising a microfluidic device that comprises:

- (a) a first elastomeric layer having a fluid flow channel within the layer, said flow channel being about 500 μm or less; and
- (b) a second elastomeric layer on top of said first elastomeric layer, said second elastomeric layer comprising a pressure channel within the second layer for controlling the flow of fluid through said fluid flow channel, and a pump and valve system coupled to the pressure channel for controlling the flow of fluid within said fluid flow channel, wherein said second elastic layer comprises a pressure channel; and
- (c) ~~a fluid flow channel within said elastomeric layer, said flow channel being about 500 μm or less; and,~~

[[~~(d)~~]] wherein the analytical system also comprises a fluid sample delivery device for delivering a sample fluid from said microfluidic device to an analytical device, said sample delivery device comprising a capillary having at least a portion thereof being located within said fluid flow channel said capillary being operatively interconnected to said analytical device for introducing said sample fluid from said fluid flow channel into said analytical device for analysis.

2-3. (Cancelled)

4. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim 1, wherein said capillary forms a hermetic seal with said flow channel.

5. (Cancelled)

6. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim 1, wherein said analytical device is selected from the group consisting of UV spectrometers, fluorescence spectrometers, IR spectrometers, gas chromatographic devices, liquid chromatographic devices, NMR devices, mass spectrometers and combinations thereof.

7. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim ~~[[6]]~~ 1, wherein said ~~sample interface means~~ fluid sample delivery device comprises a means for generating a mist from the fluid flowing through said capillary, whereby said mist is introduced to said analytical device for analysis.

8. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim 7, wherein said analytical device is a mass spectrometer.

9. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim ~~[[8]]~~ 7, wherein said means for generating a mist comprises a device for applying electrospray voltage to said capillary to generate said mist.

10. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim ~~[[9]]~~ 1, wherein ~~the tip of said capillary comprising said sample interface means is~~ comprises a tip that is tapered.

11-13. (Cancelled)

14. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim ~~[[13]]~~ 1, wherein said ~~sample interface means~~ fluid comprises generating a mist generated by using said pump and valve system.

15. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim [[13]] 1, further comprising a sample preparation chamber within said fluid flow channel.

16. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim 15, wherein said sample preparation chamber comprises a rotary fluid flow channel and a means for circulating a fluid within said rotary fluid flow channel for conducting a chemical reaction, an assay, or other sample preparations within said rotary fluid flow channel.

17. (Currently Amended) The ~~microfluidic device~~ analytical system of Claim 16, wherein said means for circulating the fluid within said rotary fluid flow channel comprises said pump and valve system.

18. (Currently Amended) An analytical apparatus for analyzing a fluid sample comprising:

- (a) an analytical device for analyzing the fluid sample; and
- (b) a microfluidic device operatively interconnected to said analytical device, wherein said microfluidic device comprises a first elastic layer comprising a fluid flow channel and a means for introducing the fluid sample into said analytical device from said fluid flow channel,

wherein said microfluidic device further comprises a second elastic layer on top of said first elastic layer, said second elastic layer comprises a pressure channel for controlling the flow of fluid through said fluid flow channel, and said microfluidic device further comprises a pump and valve system within said second elastic layer and coupled to the pressure channel for controlling the flow of fluid within said fluid flow channel.

19. (Original) The analytical apparatus of Claim 18, wherein said analytical device is selected from the group consisting of UV spectrometers, fluorescence spectrometers, IR spectrometers, gas chromatography devices, LPLC devices, HPLC devices, NMR devices, mass spectrometers and combinations thereof.

20. (Original) The analytical apparatus of Claim 19, wherein said analytical device is an electrospray ionization mass spectrometer or a nanoelectrospray mass spectrometer.

21. (Original) The analytical apparatus of Claim 20, wherein said fluid sample introducing means comprises a means for generating an ionized mist from the fluid sample.

22. (Original) The analytical apparatus of Claim 21, wherein said ionized mist generating means comprises a capillary having a distal end and a proximal end, wherein said proximal end of capillary is located within said fluid flow channel, and said distal end of capillary is interconnected to a device for applying electrospray voltage for generating the mist.

23. (Currently Amended) The analytical apparatus of Claim 22, wherein the bore diameter of said distal end of the capillary is about 100 μm or less.

24. (Currently Amended) The analytical apparatus of Claim 22, wherein said distal end of the capillary is tapered.

25-27. (Cancelled)

28. (Previously Presented) The analytical apparatus of Claim 18 further comprising a sample preparation chamber within said fluid flow channel.

29. (Original) The analytical apparatus of Claim 28, wherein said sample preparation chamber comprises a rotary fluid flow channel and a means for circulating a fluid within said rotary fluid flow channel for conducting a chemical reaction, an assay, or other sample preparations within said rotary fluid flow channel.

30. (Currently Amended) The microfluidic device of Claim 29, wherein said means mean for circulating the fluid within said rotary fluid flow channel comprises said pump and valve system.

31. (Currently Amended) A method for producing a microfluidic device comprising a means for introducing a fluid sample into an analytical device, said method comprising the steps of:

(a) producing a first elastic layer of said microfluidic device, wherein said first elastic layer comprises a fluid flow channel; and

(b) integrating a proximal end of a capillary within said fluid flow channel, wherein said step of integrating the capillary comprises the steps of:

(i.) producing a bottom portion of first elastic layer comprising a bottom portion of said fluid flow channel and a top portion of first elastic layer comprising a top portion of said fluid flow channel; and

(ii.) placing said proximal end of capillary in said bottom portion of said fluid flow channel on said bottom portion of first elastic layer and placing said top portion of the first elastic layer on top of said first portion of the first elastic layer and forming a seal between said bottom and said top portions of the first elastic layer to provide said first elastic layer comprising said fluid flow channel,

wherein a distal end of said capillary comprises said sample introducing means,

and wherein said microfluidic device further comprises a second elastic layer on top of said first elastic layer, said second elastic layer comprises a pressure channel, said microfluidic device further comprises a pump and valve system within said second elastic layer and coupled to the pressure channel for controlling the flow of fluid within said fluid flow channel.

32. (Cancelled)

33. (Original) The method of Claim 32, wherein said first elastic layer is produced by a mixture of two polymer components.

34. (Original) The method of Claim 33, wherein said bottom portion comprises an excess of one polymer component and said top portion comprises an excess of the other polymer component.

35. (Original) The method of Claim 31, wherein said distal end of capillary is interconnected to a device for applying electrospray voltage for generating a mist for introducing the fluid sample into an electrospray ionization mass spectrometer or a nanoelectrospray mass spectrometer.

36. (Original) The method of Claim 35, wherein said distal end of capillary is tapered.

37. (Currently Amended) A method for analyzing a fluid sample using an analytical device comprising the steps of:

(a) introducing said fluid sample into said analytical device through a fluid flow channel of a microfluidic device, wherein said fluid flow channel is located within a first elastic layer of said microfluidic device said microfluidic device further comprises a second elastic layer on top of said first elastic layer, said second elastic layer comprises a pressure channel, said microfluidic device further comprises a pump and valve system within said second elastic layer and coupled to the pressure channel for controlling the flow of fluid within said fluid flow channel; and

(b) analyzing said fluid sample using said analytical device.

38. (Original) The method of Claim 37, wherein said microfluidic device further comprises a sample providing means interconnected to said fluid flow channel and a sample injection site of said analytical device for introducing said fluid sample into said analytical device from said fluid flow channel.

39. (Original) The method of Claim 38, wherein said sample providing means comprises a capillary wherein a proximal end of said capillary is integrated with said fluid flow channel and the distal end of said capillary is operatively interconnected to said analytical device such that said fluid sample from said fluid flow channel is introduced into said sample injection site through said distal end of capillary.

40. (Original) The method of Claim 39, wherein said analytical device is a selected from the group consisting of UV spectrometers, fluorescence spectrometers, IR spectrometers, gas chromatographic devices, liquid chromatographic devices, NMR devices, mass spectrometers and combinations thereof.

41. (Original) The method of Claim 40, said analytical device is a mass spectrometer.

42. (Original) The method of Claim 41, wherein said sample providing means comprises generating an ionized mist from said fluid sample.

43. (Original) The method of Claim 42, wherein said ionized mist generating step comprises applying electrospray voltage to said distal end of capillary using an electrospray voltage device to generate said ionized mist.

44. (Original) The method of Claim 43, wherein the tip of said distal end of capillary is tapered.

45. (Original) The method of Claim 37, wherein said first elastic layer of microfluidic device further comprises a sample preparation chamber which is integrated with said fluid flow channel.

46-48. (Cancelled)

49. (Currently Amended) The method of Claim 37, wherein said first elastic layer comprises a sample preparation chamber integrated with said fluid flow channel, and wherein said sample preparation chamber comprises a rotary fluid flow channel and a means for circulating a fluid within said rotary fluid flow channel.

50. (Original) The method of Claim 49, wherein said mean for circulating the fluid within said rotary fluid flow channel comprises said pump and valve system.

51. (Original) The method of Claim 50 further comprising the steps of preparing said fluid sample within said sample preparation chamber.

52. (Currently Amended) The method of Claim 45, wherein said sample preparation step comprises conducting a sample preparation process within said sample preparation chamber, wherein said sample preparation process comprises:

- (i) conducting a chemical reaction;
- (ii) conducting an assay;
- (iii) degrading a peptide or protein;
- (iv) conducting a chemical analysis;
- (v) extraction of analytes from solvents;
- (vi) extraction of analytes from bodily fluids;
- (vii) concentration of sample analytes;
- (viii) affinity purification of an analyte;
- (ix) digesting a nucleic acid, carbohydrate, lipid or other molecule or mixture of molecules;

(x) separation; ~~[[and]]~~ or

(xi) mammalian, bacterial or parasite cell growth ~~(mammalian, bacterial or parasite)~~.

53. (original) The method of Claim 52, wherein said sample preparation step comprises conducting a combinatorial chemistry for preparation of an array of polymers from a monomer.

54. (Original) The method of Claim 53, wherein said monomer is selected from the group consisting of nucleotides, amino acid peptides, carbohydrates, lipids, and precursors for combinatorial synthesis.

55. (Original) The method of Claim 52, wherein said sample preparation step comprises conducting a receptor or an enzyme binding assay.

56. (Original) The method of Claim 52, wherein said sample preparation step comprises conducting binding of a target molecule to an array of oligonucleotides, peptides, proteins, oligosaccharides, or and small molecules.

57. (Original) The method of Claim 52, wherein said sample preparation step comprises conducting an enzymatic degradation of proteins, peptides, oligonucleotides, or carbohydrates, lipids, small molecules, or mixtures thereof.

58. (Currently Amended) The method of Claim ~~[[45]]~~ 37, wherein said microfluidic device comprises a plurality of sample preparation chambers which are integrated with said fluid flow channel ~~chamber~~.

59. (Currently Amended) The method of Claim 58, wherein each fluid sample from said plurality of sample preparation chambers ~~chamber~~ is independently analyzed by said analytical device.

**This Page is Inserted by IFW Indexing and Scanning
Operations and is not part of the Official Record**

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

- ☐ BLACK BORDERS
- ☐ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
- ☐ FADED TEXT OR DRAWING
- ☐ BLURRED OR ILLEGIBLE TEXT OR DRAWING
- ☐ SKEWED/SLANTED IMAGES
- ☐ COLOR OR BLACK AND WHITE PHOTOGRAPHS
- ☐ GRAY SCALE DOCUMENTS
- ☒ LINES OR MARKS ON ORIGINAL DOCUMENT
- ☐ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY
- ☐ OTHER: _____

IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.